

Message

From: Schlosser, Paul [/O=EXCHANGELABS/OU=EXCHANGE ADMINISTRATIVE GROUP (FYDIBOHF23SPDLT)/CN=RECIPIENTS/CN=121CF759D94E4F08AFDE0CEB646E711B-SCHLOSSER, PAUL]
Sent: 8/6/2021 6:31:32 PM
To: Prasad, Bidya [Prasad.Bidya@epa.gov]
Subject: RE: Model of fate of chloroprene oxidation products
Attachments: Exhibit A - Supp Mat F - Reactive Metabolite Modeling 07152021.pdf

Sure... this isn't so much a PBPK model as a cellular fate model. Attached.

-Paul

From: Prasad, Bidya <Prasad.Bidya@epa.gov>
Sent: Friday, August 6, 2021 2:15 PM
To: Schlosser, Paul <Schlosser.Paul@epa.gov>
Subject: Re: Model of fate of chloroprene oxidation products

I am by no means an expert, but I'd be interested in seeing the document to learn about what sort of metabolic activity is predicted through a PBPK model. Would you be able to send over a copy for me?

Bidya

From: Schlosser, Paul <Schlosser.Paul@epa.gov>
Sent: Friday, August 6, 2021 1:36 PM
To: Bernstein, Amanda <bernstein.amanda@epa.gov>; Brinkerhoff, Chris <Brinkerhoff.Chris@epa.gov>; Choi, Kyoungju <Choi.Kyoungju@epa.gov>; Dzierlenga, Michael <Dzierlenga.Michael@epa.gov>; El-Masri, Hisham <El-Masri.Hisham@epa.gov>; Jarabek, Annie <Jarabek.Annie@epa.gov>; Kapraun, Dustin <Kapraun.Dustin@epa.gov>; Kenyon, Elaina <Kenyon.Elaina@epa.gov>; Lin, Yu-Sheng <Lin.Yu-Sheng@epa.gov>; Morozov, Viktor <Morozov.Viktor@epa.gov>; Phillips, Martin B. <phillips.martinb@epa.gov>; Prasad, Bidya <Prasad.Bidya@epa.gov>; Sasso, Alan <Sasso.Alan@epa.gov>; Simmons, Jane <Simmons.Jane@epa.gov>; Tan, Cecilia <Tan.Cecilia@epa.gov>; White, Paul <White.Paul@epa.gov>; Zurlinden, Todd <zurlinden.todd@epa.gov>
Subject: Model of fate of chloroprene oxidation products

PKWG colleagues,

Ramboll scientists have previously developed a PBPK model for chloroprene in mice, rats, and humans, that has been under review by the EPA. In response to comments from external peer reviewers, Ramboll recently developed a provisional model to describe the fate of the two primary oxidative metabolites of CP, 1-CEO and 2-CEO. I am reading through the document now and it contains a fair level of detail regarding the chemistry, predicted fate of the metabolites.

The document will be sent back to the external panel that first reviewed the PBPK model (along with those revisions), but the panel was not formed with such a detailed chemical model in mind, and only a subset of initial reviewers are available for this follow-up review. We are seeking to add someone with in vitro metabolism expertise who has a chemistry background.

But given that, it would be good to have a couple of other internal EPA reviewers go over the analysis. If you're interested, let me know and I'll send a copy.

Thanks,
-Paul

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Paul M. Schlosser, Ph.D.

CPHEA, U.S. EPA

T [Ex. 6 Personal Privacy (PP)] e)

E: schlosser.paul@epa.gov